Docket No.: PRD-0032
App No.: Not Yet Assigned Filed: Herewith

Title: DNAs ENCODING MAMMALIAN HISTAMINE RECEPTOR OF THE H4 SUBTYPE

Inventors: Lovenberg et al. Attorney: Felicity E. Groth

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Figure 1: Nucleic Acid coding sequence human histamine H4

ATGCCAGATACTAATAGCACAATCAATTTATCACTAAGCACTCGTGTTACTTTAGCATTT TTTATGTCCTTAGTAGCTTTTGCTATAATGCTAGGAAATGCTTTGGTCATTTTAGCTTTTG ACTTCTTTGTGGGTGTGATCTCCATTCCTTTGTACATCCCTCACACGCTGTTCGAATGGG ATTTTGGAAAGGAAATCTGTGTATTTTGGCTCACTACTGACTATCTGTTATGTACAGCAT CTGTATATAACATTGTCCTCATCAGCTATGATCGATACCTGTCAGTCTCAAATGCTGTGT CTTATAGAACTCAACATACTGGGGTCTTGAAGATTGTTACTCTGATGGTGGCCGTTTGG GTGCTGGCCTTCTTAGTGAATGGGCCAATGATTCTAGTTTCAGAGTCTTGGAAGGATGA AGGTAGTGAATGTGAACCTGGATTTTTTTCGGAATGGTACATCCTTGCCATCACATCATT CTTCCAACATCTGTGGACACTCATTCAGAGGTAGACTATCTTCAAGGAGATCTCTTTCTG CATCGACAGAAGTTCCTGCATCCTTTCATTCAGAGAGACAGAGGAGAAAGAGTAGTCTC ATGTTTTCCTCAAGAACCAAGATGAATAGCAATACAATTGCTTCCAAAATGGGTTCCTT CTCCCAATCAGATTCTGTAGCTCTTCACCAAAGGGAACATGTTGAACTGCTTAGAGCCA GGAGATTAGCCAAGTCACTGGCCATTCTCTTAGGGGTTTTTGCTGTTTTGCTGGGCTCCAT ATTCTCTGTTCACAATTGTCCTTTCATTTTATTCCTCAGCAACAGGTCCTAAATCAGTTTG GTATAGAATTGCATTTTGGCTTCAGTGGTTCAATTCCTTTTGTCAATCCTCTTTTGTATCCA TTGTGTCACAAGCGCTTTCAAAAGGCTTTCTTGAAAATATTTTGTATAAAAAAGCAACC TCTACCATCACAACACAGTCGGTCAGTATCTTCTTAA

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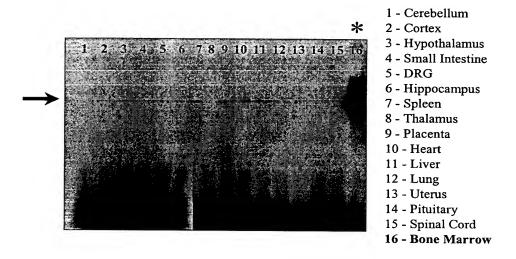
Figure 2: Amino acid sequence human histamine H4 receptor

Met Pro Asp Thr Asn Ser Thr Ile Asn Leu Ser Leu Ser Thr Arg Val Thr Leu Ala Phe Phe Met Ser Leu Val Ala Phe Ala Ile Met Leu Gly Asn Ala Leu Val Ile Leu Ala Phe Val Val Asp Lys Asn Leu Arg His Arg Ser Ser Tyr Phe Phe Leu Asn Leu Ala Ile Ser Asp Phe Phe Val Gly Val Ile Ser Ile Pro Leu Tyr Ile Pro His Thr Leu Phe Glu Trp Asp Phe Gly Lys Glu Ile Cys Val Phe Trp Leu Thr Thr Asp Tyr Leu Leu Cys Thr Ala Ser Val Tyr Asn Ile Val Leu Ile Ser Tyr Asp Arg Tyr Leu Ser Val Ser Asn Ala Val Ser Tyr Arg ThrGln His Thr Gly Val Leu Lys Ile Val Thr Leu Met Val Ala Val Trp Val Leu Ala Phe Leu Val Asn Gly Pro Met Ile Leu Val Ser Glu Ser Trp Lys Asp Glu Gly Ser Glu Cys Glu Pro Gly Phe Phe Ser Glu Trp Tyr Ile Leu Ala Ile Thr Ser Phe Leu Glu Phe Val Ile Pro Val Ile Leu Val Ala Tyr Phe Asn Met Asn Ile Tyr Trp Ser Leu Trp Lys Arg Asp His Leu Ser Arg Cys Gln Ser His Pro Gly Leu Thr Ala Val Ser Ser Asn Ile Cys Gly His Ser Phe Arg Gly Arg Leu Ser Ser Arg Arg Ser Leu Ser Ala Ser Thr Glu Val Pro Ala Ser Phe His Ser Glu Arg Gln Arg Arg Lys Ser Ser Leu Met Phe Ser Ser Arg Thr Lys Met Asn Ser Asn Thr Ile Ala Ser Lys Met Gly Ser Phe Ser Gln Ser Asp Ser Val Ala Leu His Gln Arg Glu His Val Glu Leu Leu Arg Ala Arg Arg Leu Ala Lys Ser Leu Ala Ile Leu Leu Gly Val Phe Ala Val Cys Trp Ala Pro Tyr Ser Leu Phe Thr Ile Val Leu Ser Phe Tyr Ser Ser Ala Thr Gly Pro Lys Ser Val Trp Tyr Arg Ile Ala Phe Trp Leu Gln Trp Phe Asn Ser Phe Val Asn Pro Leu Leu Tyr Pro Leu Cys His Lys Arg Phe Gln Lys Ala Phe Leu Lys Ile Phe Cys Ile Lys Lys Gln Pro Leu Pro Ser Gln His Ser Arg Ser Val Ser Ser

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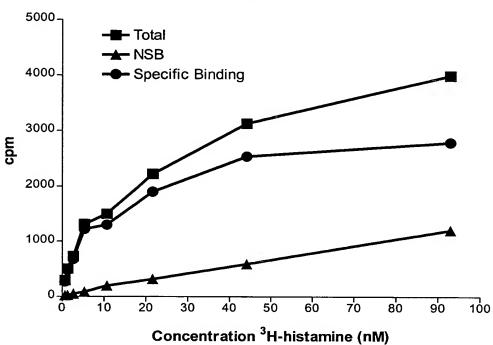
Figure 3: Tissue Distribution of human histamine H4 receptor



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Figure 4: Binding of [3H]-histamine to pH4R expressing COS7 cells.





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FIGURE 5A

Mouse H4 DNA coding region (SEQ.ID.NO.:5)

 ${\tt ATGTCGGAGTCTAACAGTACTGGCATCTTGCCACCAGCTGCTCAGGTCCCCTTGGCATTTTTAATGT}$ CTTCATTTGCCTTTGCTATAATGGTAGGCAATGCTGTGGTCATCTTAGCCTTTGTGGTGGACAGAAA CCTTAGACATCGAAGTAATTATTTTTTTTTTAATTTGGCTATTTCTGACTTCCTCGTGGGTTTGATT TCCATTCCTCTGTACATCCCTCACGTGTTGTTTAACTGGAATTTTGGAAGTGGAATCTGCATGTTTT GGCTCATTACTGACTATCTTTTGTGCACCGCATCTGTCTACAATATTGTCCTCATTAGCTACGATCG ATACCAGTCAGTTTCAAATGCTGTGTCTTATAGGGCTCAACACACTGGCATCATGAAGATTGTTGCT ${\tt CAAATGGTGGCTGTTTGGATACTGGCTTTCTTGGTAAATGGCCCGATGATTCTTGGCTTCAGATTCTT}$ GGAAGAACAGCACGAACACAAAGGACTGTGAGCCTGGCTTTGTTACAGAGTGGTACATCCTCACCAT TACAATGCTCTTGGAATTCCTGCTTCCTGTCATCTCTGTGGCTTATTTCAATGTACAGATTTACTGG AGCCTGTGGAAGCGTAGGGCTCTCAGTAGGTGCCCTAGCCATGCTGGATTCTCCACTACCTCTTCCA GTGCTTCAGGACACTTACACAGAGCTGGGGTGGCTTGCAGGACAAGTAATCCTGGATTGAAGGAATC AGCTGCATCTCGTCACTCAGAAAGTCCTCGAAGAAAGACCACCATCCTGGTGTCCTTAAGGACTCAC ATGAACAGCAGTATCACTGCCTTCAAAGTGGGTTCCTTCTGGCGATCGGAAAGTGCAGCGCTTCGCC AAAGGGAGTACGCAGAGCTTCTCAGAGGCAGGAAGCTAGCCAGGTCACTGGCCATCCTTCTGAGCGC TTTTGCCATTTGCTGGGCTCCATACTGTCTGTTCACAATTGTCCTTTCAACTTACCCCAGAACGGAA CGCCCAAATCGGTGTGCTACAGCATTGCCTTCTGGCTGCAATGGTTCAATTCGTTTGTTAATCCCT TTCTGTACCCTTTGTGTCACAGGCGTTTCCAGAAGGCTTTCTGGAAGATACTTTGTGTGACAAAGCA ACCAGCGCTGTCACAGAACCAGTCAGTATCTTCTTGA

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FIGURE 5B

Rat H4 DNA coding region (SEQ.ID.NO.:6)

ATGTCGGAGTCTAACGCCACTGACGTCTTGCCACTGACTCCACTCAAGTCCCCTTGGCATTTTTAATGT CCTTAGACATCGAAGTAATTATTTTTTCTTAATTTGGCTATTTCTGACTTCTTCGTGGGTGTCATC TCCATTCCTCTGTACATCCCTCACACGCTGTTTAACTGGAATTTTGGAAGTGGAATCTGCATGTTTT GGCTCATTACTGACTATCTTTTGTGCACAGCATCCGTCTACAGTATTGTCCTCATTAGCTACGATCG ATACCAGTCAGTTTCAAACGCTGTGCGTTATAGAGCACAGCACACTGGCATCCTGAAAATTGTTGCT CAAATGGTGGCTGTTTGGATACTGGCTTTCTTGGTCAATGGCCCAATGATTCTGGCTTCGGATTCTT GGAAGAACAGCACCAACACAGAGGAGTGCGAGCCTGGCTTTGTTACTGAGTGGTACATCCTCGCCAT TACAGCATTCTTGGAATTCCTGCTCCCTGTCTCCTTGGTGGTCTATTTCAGTGTACAGATTTACTGG AGCCTGTGGAAGCGTGGGAGTCTCAGTAGGTGCCCTAGCCACGCTGGATTCATCGCTACCTCTTCCA GGGGCACTGGACACTCACGCAGAACTGGGTTGGCTTGTAGGACAAGTCTTCCTGGATTAAAGGAACC ATGAGCGGTAGTATCATCGCCTTCAAAGTGGGTTCCTTCTGCCGATCAGAAAGCCCAGTGCTTCACC AGAGAGAGCACGTGGAGCTTCTCAGAGGCAGGAAGCTAGCCAGGTCGCTAGCTGTCCTCCTGAGTGC $\tt TTTTGCCATTTGCTGGGCTCCGTATTGCCTGTTCACAATTGTTCTTTCAACTTATCGCAGAGGGGAG$ CGCCCCAAATCGATTTGGTACAGCATAGCCTTTTGGCTACAGTGGTTCAATTCACTTATTAATCCCT TTCTATACCCTTTGTGCCACAGACGTTTCCAGAAGGCTTTCTGGAAGATACTCTGTGTGACAAAGCA ACCAGCACCTTCACAGACCCAGTCAGTATCTTCTTGA

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FIGURE 5C

Guinea Pig H4 DNA coding sequence (SEQ.ID.NO.:7)

ATGTTGGCAAATAACAGTACAATCGCCTTAACATCAATTAAAATTTCTTTGACATTTTA ATGTCTTTACTAGCTATTGCTATAATGTTAGGCAATGTCGTGGTCATTTTAGCTTTTATTG TCTTTGTGGGTGCAATTGCAATTCCTCTGTACATACCTTCCTCGCTGACTTACTGGACTT $\tt CTGGAAAGCAAGCTTGTGTATTTTGGCTCATTACTGACTATCTTTTATGtACAGCATCTGT$ TAGAGCTCAGCACTCTGGCACCTGGAAAATTGCTACTCAGATGGTGGCTGTTTGGATAT TCTCCTTCATGACAAATGGGCCGATGATTCTGATTTCAGACTCTTGGCAGAATAGCACT ACAGAATGTGAACCTGGATTTTTAAAAAAGTGGTACTTTGCTCTCCCTACATCATTATTG GAATTCCTGATCCCCATCTTGTTAGTTGCTTATTTCAGCGCCCATATTTACTGGAGCCTG TGGAAGCGAGAGAAACTGAGCAGGTGCCTCAGCCACCCTGTACTCCCCTCTGACTCTTC CAGCAGTGACCACGGACACTCCTGCAGACAGGACCCCGATTCAAGGGCGACTCTGCCA GCACGGAAAGAACAACTGCCTCTCTTGGTTCAGACAAGTCACGGAGAAAGAGCAGTC TCTTGTTTTCCATAAGAGCCTACAAGAACAGCAATGTGATCGCTTCCAAAATGGGCTTC CTCTCCCACTCAGATTCCCTGGCTCTTCAGCAAAGGGAACATATCGAACTTTTCAGAGC CAGGAAATTAGCCAAGTCACTGGCCATACTCTTAGCAGCTTTTGCCATTTGCTGGGCTCC ATATTCACTGACTACAGTTATCTACTCATTTTTCCTGAAAGGAACTTGACTAAATCAAC CTGGTACCATACTGCCTTTTGGCTCCAGTGGTTCAATTCCTTTGTTAATCCCTTTTTGTAT CCATTGTGTCACAAACGTTTTCAGAAGGCTTTCCTGAAAATACTTCCTGTGAGAAGGCA ATCCACGCCACACACACCGCTCAATATCCACTTGA

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FIGURE 6A

Mouse histamine H4 receptor protein sequence (SEQ.IN.NO.:8)

MSESNSTGILPPAAQVPLAFLMSSFAFAIMVGNAVVILAFVVDRNLRHRSNYFFLNLAISDFL VGLISIPLYIPHVLFNWNFGSGICMFWLITDYLLCTASVYNIVLISYDRYQSVSNAVSYRAQH TGIMKIVAQMVAVWILAFLVNGPMILASDSWKNSTNTKDCEPGFVTEWYILTITMLLEFLLP VISVAYFNVQIYWSLWKRRALSRCPSHAGFSTTSSSASGHLHRAGVACRTSNPGLKESAASR HSESPRRKSSILVSLRTHMNSSITAFKVGSFWRSESAALRQREYAELLRGRKLARSLAILLSA FAICWAPYCLFTIVLSTYPRTERPKSVWYSIAFWLQWFNSFVNPFLYPLCHRRFQKAFWKIL **CVTKQPALSQNQSVSS**

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FIGURE 6B

Rat histamine H4 receptor protein sequence (SEQ.ID.NO.:9)

MSESNGTDVLPLTAQVPLAFLMSLLAFAITIGNAVVILAFVADRNLRHRSNYFFLNLAISDFF VGVISIPLYIPHTLFNWNFGSGICMFWLITDYLLCTASVYSIVLISYDRYQSVSNAVRYRAQH TGILKIVAQMVAVWILAFLVNGPMILASDSWKNSTNTEECEPGFVTEWYILAITAFLEFLLPV SLVVYFSVQIYWSLWKRGSLSRCPSHAGFIATSSRGTGHSRRTGLACRTSLPGLKEPAASLH SESPRGKSSLLVSLRTHMSGSIIAFKVGSFCRSESPVLHQREHVELLRGRKLARSLAVLLSAF AICWAPYCLFTIVLSTYRRGERPKSIWYSIAFWLQWFNSLINPFLYPLCHRRFQKAFWKILCV TKQPAPSQTQSVSS

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FIGURE 6C

Guinea Pig histamine H4 receptor protein sequence (389 amino acids) (SEQ.ID.NO.:10)

MLANNSTIALTSIKISLTFLMSLLAIAIMLGNVVVILAFIVDRNLRHRSNYFFLNLAIADFFVG AIAIPLYIPSSLTYWTSGKQACVFWLITDYLLCTASVYNIVLISYDRYQSVSNAVWYRAQHS GTWKIATQMVAVWIFSFMTNGPMILISDSWQNSTTECEPGFLKKWYFALPTSLLEFLIPILLV A YFSAHIYWSLWKREKLSRCLSHPVLPSDSSSSDHGHSCRQDPDSRATLPARKETTASLGSDKSRRKSSLLFSIRAYKNSNVIASKMGFLSHSDSLALQQREHIELFRARKLAKSLAILLAAFAIC WAPYSLTTVIYSFFPERNLTKSTWYHTAFWLQWFNSFVNPFLYPLCHKRFQKAFLKILPVRR **QSTPPHNRSIST**

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TCCTTCTGGCGATCGGAAAGTGCAGCGCTTCGCCAAAGGGAGTACGCAGAGCTTCTCAGAGGCAAGCTAGCCAGGTCACTGGCCATCCTTCTGAGCGCTTTTGCCATTTTG

Nucleotide Sequence Comparison of Human (SEQ ID NO:1), Guinea Pig (SEQ ID NO:7), Mouse (SEQ ID NO:5), and Rat H4 (SEQ ID NO:6)

Figure 7A

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480
                                                                                                                                                                                                                       TGGTATAGAGCTCAGCACTCTGGGAAAATTGCTACTCAGATGGTGGCTGTTTGGATATTCTCCTTCATGACAAATGGGCCGATGATTCTGATATTCAGACTCTTGGCAGAATAGC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   900
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         TCTTATAGGGCTCAACACTGGCATCATGAAGATTGTTGCTCAAATGGTGGCTGTTTGGATACTGGCTTTCTTGGTAAATGGCCCGATGATGTTCTGGCTTCAGATTCTTGGAAGAACAGC 480
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      837
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                                                                                                                                          CGTTATAGAGCACACACTGGCATCCTGAAAATTGTTGCTCAAATGGTGGCTGTTTGGATACTGGCTTTCTTGGTCAATGGCCCAATGATTCTGGCATTCTTGGAAGAACAGC
Human: ATGCCAGATACTAATAGCACAATCAATTATCACTAAGCACTCGTGTTACTTTAGGTGTTTTTTATGTCCTTAGTAGCTTTTGCTAAATGCTTTGGTAGGAAATG
                                                                       Guinea Pig: ATGTTGGCAAATAACAGTACAATGGCCTTAACA---TCAATTAAAATTTCTTTGACATTTTTAATGTCTTTACTAGCTATTGGTATGTTAAGGCAATGTCGTGGTCATTTAGCTTTTAGCTTA
                                                                                                                                                                                                                                                                                                                                                 AATTITGGAAGIGGAATCIGCAIGTITITGGCICATIACIGACIAICITITIGIGCACAGCAICCGICIACAGIAIITGTCTCAGIAIGCGAGAGAGATAGCAGICAGICAGICAGICAGICAGICAGITICAAAACGCIGIG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ACCAACACAGAGGAGGCGGGGCTTGTTACTGAGTGGTACATCCTCGCCATTACAGCATTCTTGGAATTCCTGCTCCCTGTCTTGGTGGTGTCTATTTCAGTGTACAGATTTC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Guinea Pig: ACT---ACA---GAATGTGAACCTGGATTTTTAAAAAAGTGGTACTTTGCTCTCCCTACATCATTATTGGAATTCCTGATCCTGATCTTGTTAGTTGCTTATTTCAGCGCCCATATTTACA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 TGGAGCCTGTGGAAGGTTAGGTGTTCAGTAGGTGCCCTAGCCATGCTGGATTC---TCCACTACCTCTTCCAGTGCTTCAGGACACTTACACAGAGCTGGGGTGGCTTGCAGGAGT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   GATTTTGGAAAGGAAATCTGTGTGTATTTTGGCTCACTACTGACTATCTGTTATGTACAGCATCTGTATATAAACATTGTCCTCATCAGCTATGATCGATACCTGTCAGTCTGAAATGCTGT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                GGT---AGT---GAATGTGAACCTGGATTTTTTCGGAATGGTACATCCTTGCCATCACATCTTGGAATTCGTGATCCCAGTCATCTTAGTCGCTTATTTCAACATGAATATTTAT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                CTTCCTGGATTAAAGGAACCAGCCGCATCCCTTCATTCAGAAAGTCCACGAGGAAAGAGCAGTCTCCTGGTGTCCTTAAAGGACTCACATGAGGGTAGTATCATCACTTCAAAGTGGGT
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         TGGAGCCTGTGGAAAGGGAGAAACTGAGCAGGTGCCTCAGCCACCCTGTACTCCCCTCTGACTCTTCCAGCAGTGACCAGGACACTCCTGCAGACAGGACAGGACCGGATTCAAGGGGGACA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    IGGAGCCTGTGGGAAGCGTGGGAGTCTCAGTAGGTGCCCTAGCCACGCTGGATTC---ATCGCTACCTCTTCCAGGGGCACTGGACTCACGCAGAACTGGGTTGGCTTGTAGGACA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 CTGCCAGCACGGAAAGAAACAACTGCCTCTTGGTTCAGACAAGTCACGGAGAAAGAGCAGTCTCTTGTTTTCCATAAGAGGCCTACAAGAAGCAGTGTGGATGTGATCGCTTCCAAAAATGGGC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   TCCTTCTCCCAATCAGATTCTGTAGCTCTTCACCAAAGGGAACATGTTGAACTGCTTAGAGCCAGGAGATTAGCCAAGTCACTGGCCATTCTCTTAGGGGTTTTGCTGTTTGCTGGCCA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        aatcctggattgaaggaatcagctgcatctcgtcactcagaaagtcctcgaagaaggcagcatcctggtgtccttaaggactcacatgaacagcagtatcactgccttcaaagtggg
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             TICCTCTCCCCACTGABATTCCCTGGCTCTTCAGCAAAGGGAACATATCGAACTTTTCAGAGCCAGGAAATTAGCCAAGTCACTGGCCATACTCTTAGCAGCTTTTGCTGGCTCT
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Figure 7B

TTG 1071 TTG 1071 CTG 1077	1173 1170 1176 1176		
GTGGTTCAATTCCTTTGTCAATCCTCTT GTGGTTCAATTCCTTTGTTAATCCCTTT ATGGTTCAATTCGTTTGTTAATCCCTTT GTGGTTCAATTCACTTATTAATCCCTTT	AGTATCTTTAA AATATCCACTTGA AGTATCTTCTTGA AGTATCTTCTTGA		
HTTGGTATAGAATTGCATTTTGGCTTCA NCTGGTACCATACTGCCTTTTGGCTCCA HTTGGTACAGCATTGCCTTTTGGCTGCA NTTGGTACAGCATAGCCTTTTGGCTACA	AAGGCTTTCTTGAAAATATTTTGTATAAAAAGCAACCTCTACCATCACAACACAGTCGGTCAGTATTCTTGTAA AAGGCTTTCCTGAAAATACTTCCTGTGAGAAGGCAATCCACGCCCACACAGAACCGCTCAATATCCACTTGA AAGGCTTTCTGGAAGATACTTTGTGTGACAAAGCAACCAGCGCTGTACAGAACCAGTCAGTATCTTCTTGA		
ATTCCTCAGCAACAGGTCCTAAATCAG TTCCTGAAAGGAACTTGACTAAATCAA ACCCCAGAACGGAACG	TCTTGAAAATATTTTGTATAAAAAGG TCCTGAAAATACTTCCTGTGAGAGGG TCTGGAAGATACTTTGTGTGACAAAGCJ TCTGGAAGATACTCTGTGTGACAAAGCJ		
Human: CCATATTCTCTGTCACAATTGTCCTTTCATTTATCCTCAGCAACAGGTCCTAAATCAGTTTGGTATAGAATTGGCTTTGGGGTTCAATTCCTTTGTCAATCCTTTTTG 1071 a Pig: CCATATTCACTGACTACAGTTATCTACTCATTTTTCTGAAGAACTTGACTAAATCAACCTGGTACCATACTGGCTTTTGGCTCCAGTGGTTCAATTCCTTTGTTAATCCCTTTTTG 1071 Mouse: CCATACTGTTCACAATTGTCCTTTCAACTTACCCCAGAACGGAACGCCCCAAATCGTACAGCATGGCTTCTGGCTGCTACAATTGGTTAATCCCTTTCTG 1077 Rat: CCGTATTGCCTGTTCACAATTGTTCTTTCAACTTATCGCAGAGGGGAGCGCCCCAAATCGATTGGTACAGCATTGGCTATTGGCTTATTGGCTAGCTTATTGGCTAATTCACTTATTAATCCCTTTTTTTA 1077	TATCCATTGTGTCACAAGCGCTTTCAAAAGGCTTTCTTGAAAATATTTTGTATAAAAGCAACCTCTACCATCACAACACGCGGCTGGTATCTTTTAA TATCCATTGTGTCACAAACGTTTTCAGAAGGCTTTCCTGAAAATACTTCCTGTGAGAAGGCAATCCACGCCACCACACAACCGCTCAATATCCACTTGA TACCCTTTGTGTCACAGGCGTTTCCAGAAGGCTTTCTGGAAGATACTTTGTGTGTG		
Human: CCJ Guinea Pig: CCJ Mouse: CCJ Rat: CCC	Human: TAT Guinea Pig: TAT Mouse: TAC Rat: TAC		

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390 389 391 391

ARRLAKSLAILLGVFAVCWAPYSLFTIVLSFYSSATGPKSVWYRIAFWLQWFNSFVNPLLYPLCHKRFQKAFLKIFCIKKQPLPSQHSRSVSS ARKLAKSLAILLAAFAICWAPYSLTTVIYSFFPERNLTKSTWYHTAFWLQWFNSFVNPFLYPLCHKRFQKAFLKILPVRRQSTPP-HNRSIST GRKLARSLAILLSAFAICWAPYCLFTIVLSTYPRTERPKSVWYSIAFWLQWFNSFVNPFLYPLCHRRFQKAFWKILCVTKQPALSQ-NQSVSS GRKLARSLAVLLSAFAICWAPYCLFTIVLSTYRRGERPKSIWYSIAFWLQWFNSLINPFLYPLCHRRFQKAFWKILCVTKQPAPSQ-TQSVSS

Human: Guinea Pig: Mouse: Rat: . .KS.WY:.AFWLQWFNS::NP:LYPLCH:RFQKAF.KI: :::Q: .:

.R:LA:SLA:LL:.FA:CWAPY:L T:: S

Consensus:

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Amino Acid sequence Comparison of Human (SEQ ID NO:2), Guinea Pig (SEQ ID NO:10), Mouse (SEQ ID NO:8) and Rat H4 (SEQ ID NO:9) Figure 8

100 99 100 100	198 197 200 200	297 297 299 299
Human: MPDT <u>N</u> STINLSLSTRVTLAFFMSLVAFAIMLGNALVILAFVVDKNLRHRSSYFFINLAISDFFVGVISIPLYIPHTLFEWDFGKEICVFWLTTDYLLCTA Guinea Pig: MLA <u>NN</u> STIALT-SIKISLTFLMSLAIAIMLGNVVVILAFIVDRNLRHRSNYFFLNLAIADFFVGAIAIPLYIPSSLTYWTSGKQACVFWLITDYLLCTA Mouse: MSES <u>N</u> STGILPPAAQVPLAFLMSSFAFAIMVGNAVVILAFVVDRNLRHRSNYFFLNLAISDFLVGLISIPLYIPHVLFNWNFGSGICMFWLITDYLLCTA Rat: MSES <u>N</u> GTDVLPLTAQVPLAFLMSLAFAITIGNAVVILAFVADRNLRHRSNYFFLNLAISDFFVGVISIPLYIPHTLFNWNFGSGICMFWLITDYLLCTA Consensus: MN.T .L: ::::L:F:MSL:A:AIM:GN.:VILAF::D:NLRHRS:YFFINLAI:DF:VG.I:IPLXIP.:L TM3 TM3 TM3	Human: SVYNIVLISYDRYLSVSNAVSYRTQHTGVLKIVTLMVAVWVLAFLVNGPMILVSESWKDEG-S-ECEPGFFSEWYILAITSFLEFVIPVILVAYFSAHIY Guinea Pig: SVYNIVLISYDRYQSVSNAVWYRAQHSGTWKIATQMVAVWIFSFMTNGPMILISDSWQNST-T-ECEPGFLKKWYFALPTSLLEFLIPILLVAYFSAHIY Mouse: SVYNIVLISYDRYQSVSNAVSYRAQHTGIMKIVAQMVAVWILAFLVNGPMILASDSWKNSTNTKDCEPGFVTEWYILTITMLLEFLLPVISVAYFNVQIY Rat: SVYSIVLISYDRYQSVSNAVRYRAQHTGILKIVAQMVAVWILAFLVNGPMILASDSWKNSTNTEECEPGFVTEWYILAITAFLEFLLPVSLVVYFSVQIY Consensus: SVXNIVLISYDRY SVSNAV YR:QH:G. KI MVAVW:::F:.NGPMIL:S:SW:::CEPGF:WY:::T.:LEF::P::VAYF:.:IY TM5	Human: WSLWKRDHLSRCQSHPGL-TAVSSNICGHSFRGRLSSRRSLSASTEVPASFHSERQRRKSSLMFSSRTKMNSNTIASKMGSFSQSDSVALHQREHVELLR Guinea Pig: WSLWKREKLSRCLSHPVLPSDSSSSDHGHSCRQDPDSRATLPARKETTASLGSDKSRRKSSLLFSIRAYKNSNVIASKMGFLSHSDSLALQQREHIELFR Mouse: WSLWKRRALSRCPSHAGF-STTSSSASGHLHRAGVACRTSNPGLKESAASRHSESPRRKSSILVSLRTHMNSSITAFKVGSFWRSESAALRQREYAELLR Rat: WSLWKRGSLSRCPSHAGF-IATSSRGTGHSRRTGLACRTSLPGLKEPAASLHSESPRGKSSLLVSLRTHMSGSIIAFKVGSFCRSESPVLHQIEHVELLR Consensus: WSLWKR: LSRC SH. : SS. GH. R R : :
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